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## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

| Application<br>Number     | 09/461,090        |
|---------------------------|-------------------|
| Filing Date               | December 14, 1999 |
| First Named<br>Inventor   | ULLRICH et al.    |
| Group Art Unit            | 1634              |
| Examiner Name             | F. Lu             |
| Attorney Docket<br>Number | 2923-0347         |

Title of the Invention: EGF RECEPTOR TRANSACTIVATION BY G-PROTEIN-COUPLED RECEPTORS REQUIRES METALLOPROTEINASE CLEAVAGE OF Pro-HB-EGF

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## SUPPLEMENTAL AMENDMENT UNDER 37 CFR 1.121

TECH CENTER 1600/2900

Commissioner for Patents Washington, D.C. 20231

February 21, 2003

Dear Sir:

In further response to the non-final Office Action dated May 21, 2002, Applicants request entry of the amendments below and reconsideration of the application.

## In the Claims:

Please amend the claims as follows:

36.(Amended) A method for modulating a G-protein mediated signal transduction comprising:

providing a cell having a disturbed G-protein mediated signal transduction and a receptor tyrosine kinase capable of transactivation by G-protein mediated signal transduction, wherein said receptor tyrosine kinase is selected from the group consisting of EGFR and other members of the EGFR family, said receptor tyrosine kinase comprising an extracellular domain and said cell having a G-protein mediated signal transduction pathway